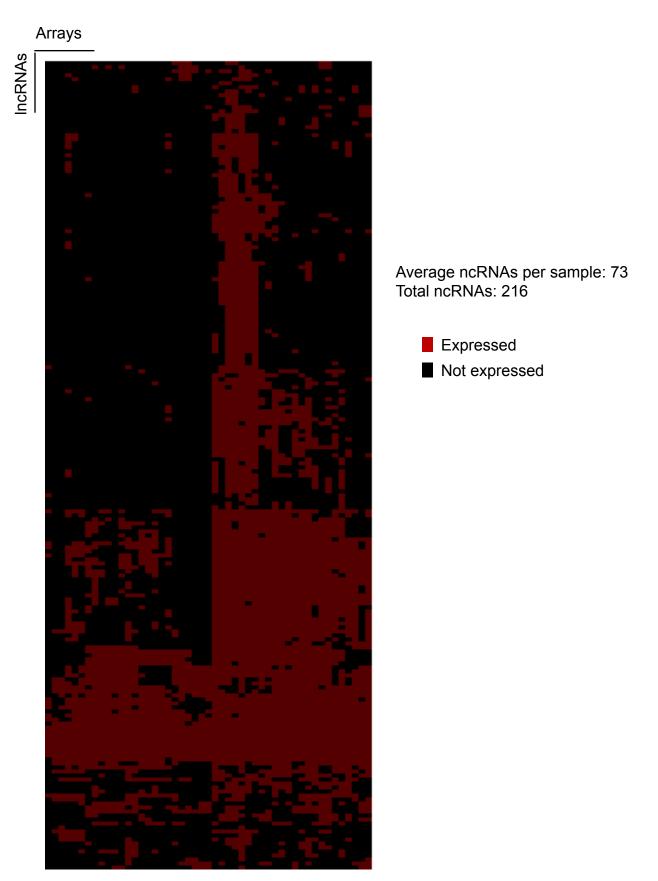
Supplementary Data:

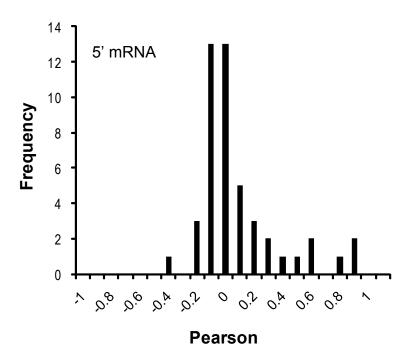
Extensive and coordinated transcription of noncoding RNAs within cell cycle promoters

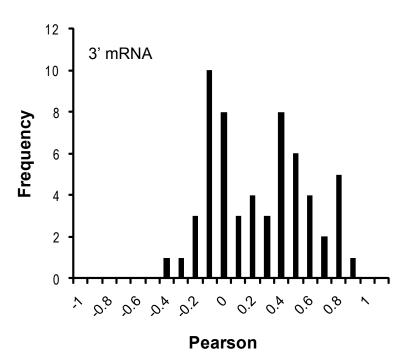
Tiffany Hung^{1,2}, Yulei Wang³, Michael F. Lin^{4,5}, Ashley K. Koegel^{1,2}, Yojiro Kotake⁶⁻⁸, Gavin Grant⁹, Hugo M. Horlings¹⁰, Nilay Shah¹¹, Christopher Umbricht¹², Pei Wang¹³, Yu Wang³, Benjamin Kong³, Anita Langerod^{14,15}, Seung K. Kim^{2,13}, Marc van de Vijver¹⁰, Saraswati Sukumar¹¹, Michael L. Whitfield⁹, Manolis Kellis^{4,5}, Yue Xiong⁶, David J. Wong^{1*}, Howard Y. Chang^{1,2*}

Supplementary Figure 1. Heatmap of lncRNAs expressed in each of the 104 different RNA tiling arrays as determined by peak calling analysis (see Methods).



Supplementary Figure 2. RT-PCR validated expression correlation between 60 lncRNAs and their nearest 3'and 5' mRNAs across 34 RNA samples.

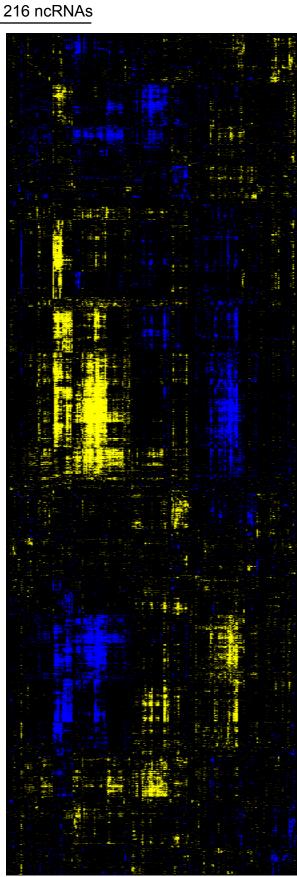




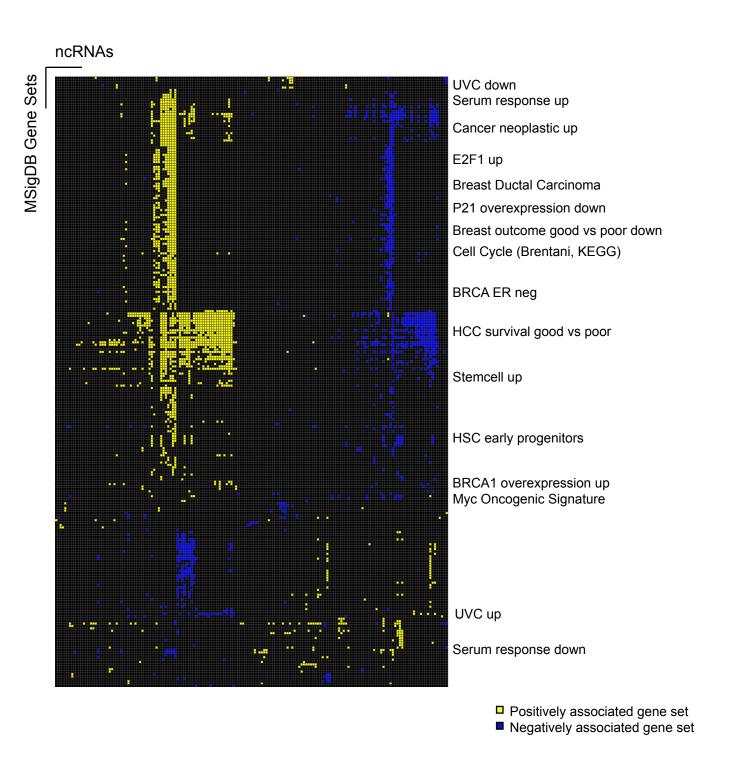
Supplementary Figure 3.

Gene sets of mRNAs positively or negatively correlated with each lncRNA as determined by pairwise Pearson correlation across 17 tiling and expression arrays.

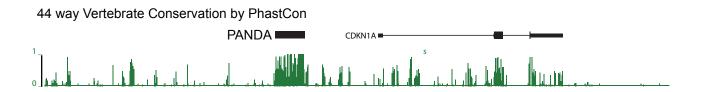
10479 mRNAs



- mRNA negatively correlated with ncRNA
- mRNA positively correlated with ncRNA
- mRNA with no correlation to ncRNA

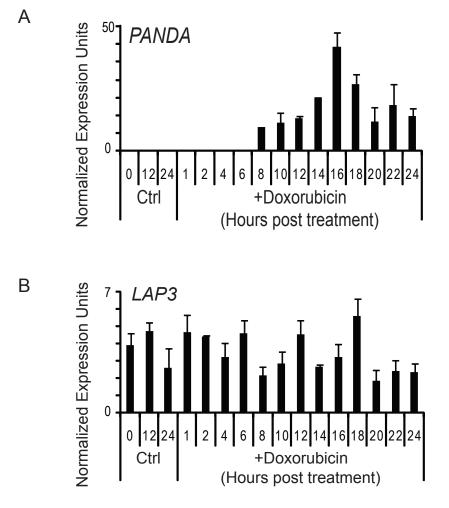


Supplementary Figure 5. PANDA is evolutionarily conserved across vertebrates as determined by 44 way Vertebrate Conservation PhastCon score.



Supplementary Figure 6.

24 hour DNA damage time course of PANDA and LAP3 expression. Human fetal lung fibroblasts (FL3) cells were treated with doxorubicin and collected at the indicated time points for RT-PCR analysis.



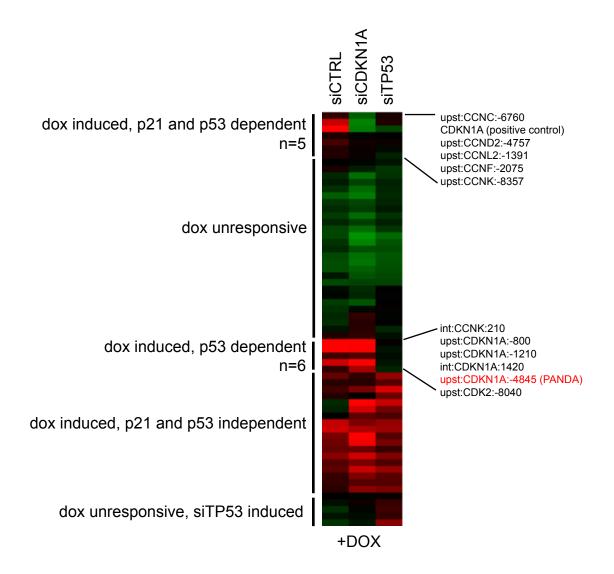
Supplementary Figure 7. RLM-RACE sequence of PANDA

A. 5'RACE sequence of PANDA

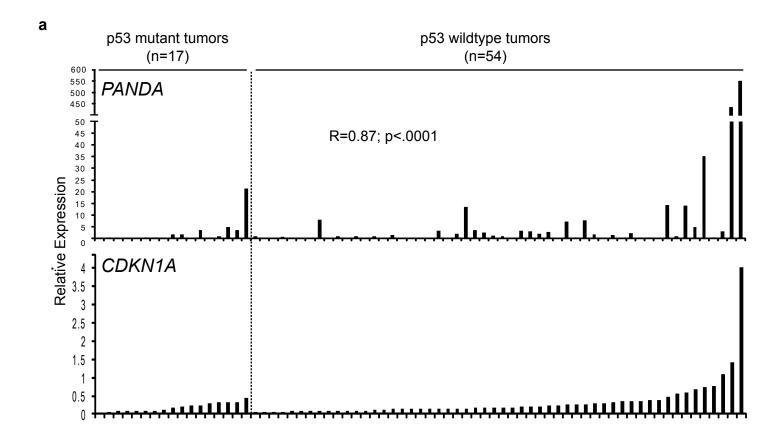
TAGATGTAGCTTTGGGGTCAGGTGCCACTGGGGTCTTTACCAATTCATCCTGGCTCTGGAACAAGCTC
TTCGAGGCCAGCATTGAAACAGGGGACCGTGTCTGGAGGATGCCTCTCTTCAAACATTGTACAAGAC
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CTGCGGCATTCCTGAAAGAATTCGT

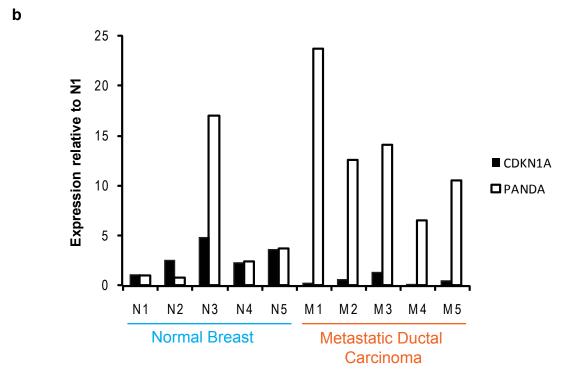
B. 3' RACE sequence of PANDA

Supplementary Figure 8. p53-dependent DNA damage induction in a subset of lncRNAs. Heatmap of lncRNA expression (as measured by RT-PCR) of human fetal lung fibroblasts (FL3) treated with doxorubicin in the presence of siCTRL, siCDKN1A, or siTP53. Red indicates induction relative to undamaged cells, green indicates repression.



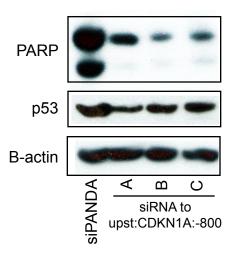
Supplementary Figure 9. PANDA expression level in tumors. (A) Expression in p53 mutant vs. p53 wildtype tumors. Human primary breast tumors were derived from the fresh-frozen tissue bank of the Netherlands Cancer Institute/Antoni van Leeuwenhoek Hospital . For mutational analysis, DNA extraction was performed as described previously1 and TP53 mutations were identified by DNA sequencing for exons 2-11 as described2. (B) Expression of PANDA in 5 normal breast tissues vs. 5 metastatic ductal carcinomas, also obtained from the same tissue depository as (A).



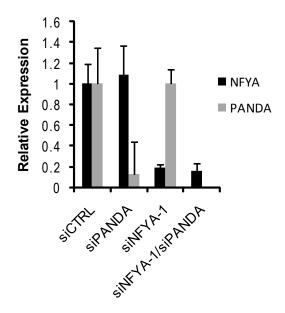


1. Horlings, H.M. et al. Integration of DNA copy number alterations and prognostic gene expression signatures in breast cancerapients. Clin Cancer Res 16, 651-63.

Supplementary Figure 10. Three independent siRNAs to upst:CDKN1A:-800 did not induce PARP cleavage in FL3 cells upon treatment with doxorubicin.



Supplementary Figure 11. Knockdown efficiency of NFYA and PANDA for Fig 7d and 7e.



Supplementary Table 1: Tiling Array design

	rable I. Illing	
<u>Name</u>		Feature Coordinates (Human March 2006 NCBI Build 36.1 hg18)
9p21 locus		21900000-22150000
CCNA1		35894632-35906659
CCNA2	4	122962330-122974342
CCNB1	5	68488668-68500750
CCNB2	15	57174611-57186627
CCNB3	X	50034275-50046275
CCNC	6	100121225-100133411
CCND1	11	69155053-69167126
CCND2	12	4243198-4255223
CCND3		42015122-42027530
CCNE1		34984740-34997400
CCNE2		95974605-95986660
CCNF		2409440-2421471
CCNG1		162787154-162799204
CCNG2		78287401-78299550
CCNH		86742441-86754592
CCNI		78214148-78226149
CCNJ		97783140-97795329
CCNJL		159697177-159709177
CCNK		99007491-99019512
CCNL1		158358577-158371176
CCNL2		1322552-1334571
CCNO		54563265-54575265
CCNT1		47395048-47407048
CCNT2		135382862-135394875
CCNY		35565959-35577959
CCNYL1		208274509-208286509
CCNYL2		42268168-42280168
CCNYL3		34105360-34117360
CDK2		54636825-54648899
CDK3		71499013-71511013
CDK4	12	56430344-56442431
CDK5		150383893-150395929
CDK5R1	17	27828217-27840681
CDK5R2	2	219522620-219534641
CDK6	7	92299148-92311148
CDK8	13	25716755-25728778
CDK9	9	129578151-129590188
CDK10	16	88270578-88282613
CDKL1	14	49930367-49942367
CDKL2	4	76772269-76784595
CDKL3	5	133728113-133740664
CDKL4	2	39308177-39320177
CDKL5	Х	18343677-18355677
CDKN1A	6	36744464-36756493
CDKN1B		12751575-12763663
CDKN1C		2861551-2873577
CDKN2C		51196148-51210203
CDKN2D		10538631-10550655
CDKN3		53923462-53935475
J	17	

CNNM1	10 101070022-101082022
CNNM2	10 104658064-104670853
CNNM3	2 96835714-96848658
CNNM4	2 96780365-96792606

Hung et al., Supplementary Table 2

Sample pair # Experimental sample

- 1 Human fetal lung fibroblasts treated with doxorubicin (200ng/ul) for 24 hours
- 2 Human fetal lung fibroblasts in low serum (0.01%)
- 3 Human fetal lung fibroblasts transduced with HPV-E7
- 4 Human fetal lung fibroblasts transduced with HPV-E6
- 5 Human fetal lung fibroblasts transduced with HRAS
- 6 HeLa synchronized by double thymidine block: 0 hr
- 7 HeLa synchronized by double thymidine block: 2 hr
- 8 HeLa synchronized by double thymidine block: 4 hr
- 9 HeLa synchronized by double thymidine block: 6 hr
- 10 HeLa synchronized by double thymidine block: 8 hr
- 11 HeLa synchronized by double thymidine block: 10 hr
- 12 HeLa synchronized by double thymidine block: 12 hr
- 13 Primary human keratinocytes transduced with p63 shRNA
- 14 Primary human keratinocytes treated with Ca2+ for 48 hours
- 15 U2OS synchronized by double thymidine block: 0 hr
- 16 U2OS synchronized by double thymidine block: 2 hr
- 17 U2OS synchronized by double thymidine block: 4 hr (A)
- 18 U2OS synchronized by double thymidine block: 4 hr (B)
- 19 U2OS synchronized by double thymidine block: 6 hr
- 20 U2OS synchronized by double thymidine block: 8 hr
- 21 U2OS synchronized by double thymidine block: 14 hr
- 22 U2OS synchronized by double thymidine block: 16 hr
- 23 Human ES (H9)
- 24 Human ES (H9)
- 25 Human ES (H9)
- 26 MCF7 cell line
- 27 Primary human keratinocytes transduced with MYC
- 28 Primary human keratinocytes transduced with MYC
- 29 Primary human keratinocytes transduced with HRAS
- 30 Primary human keratinocytes transduced with HRAS
- 31 Primary human keratinocytes transduced with E2F3
- 32 Primary human keratinocytes transduced with E2F3
- 33 Primary human keratinocytes transduced with IkB
- 34 Primary human keratinocytes transduced with IkB
- 35 Primary human keratinocytes transduced with MYC, RAS, and IkB
- 36 Primary human keratinocytes transduced with MYC, RAS, and IkB
- 37 Primary human keratinocytes transduced with E2F3, RAS, and IkB
- 38 Primary human keratinocytes transduced with E2F3, RAS, and IkB
- 39 Primary human keratinocytes transduced with SOX2, RAS, and IkB 40 Primary human keratinocytes transduced with SOX2, RAS, and IkB
- 41 MYC-RAS-IkB tumor 1
- 42 MYC-RAS-IkB tumor 2
- 43 E2F3-RAS-IkB tumor 1
- 44 E2F3-RAS-IkB tumor 2
- 45 Invasive ductal breast carcinoma P2
- 46 Invasive ductal breast carcinoma P3
- 47 Invasive ductal breast carcinoma P4
- 48 Invasive ductal breast carcinoma P5
- 49 Invasive ductal breast carcinoma P6
- 50 Invasive ductal breast carcinoma P7
- 51 Invasive ductal breast carcinoma P9
- 52 Invasive ductal breast carcinoma P10

Control sample

Human fetal lung fibroblasts untreated

Human fetal lung fibroblasts in normal serum

Human fetal lung fibroblasts transduced with vector control

Human fetal lung fibroblasts transduced with vector control

Human fetal lung fibroblasts transduced with vector control

HeLa asynchronous

Primary human keratinocytes transduced with control shRNA

Primary human keratinocytes without Ca2+ treatment

U2OS asynchronous

Human SOX17+ definitive endoderm

Human fetal pancreas day 76

Human fetal pancreas day 152

Human mammary epithelial cells

Primary human keratinocytes transduced with GFP

Primary human keratinocytes transduced with LacZ

Primary human keratinocytes transduced with GFP

Primary human keratinocytes transduced with LacZ

Primary human keratinocytes transduced with GFP

Primary human keratinocytes transduced with LacZ

Primary human keratinocytes transduced with GFP

Primary human keratinocytes transduced with LacZ

Primary human keratinocytes transduced with LacZ, RAS, and IkB

Primary human keratinocytes transduced with GFP, RAS, and IkB

Primary human keratinocytes transduced with LacZ, RAS, and IkB

Primary human keratinocytes transduced with GFP, RAS, and IkB

Primary human keratinocytes transduced with LacZ, RAS, and IkB

Primary human keratinocytes transduced with GFP, RAS, and IkB

GFP-RAS-IkB tumor pool

GFP-RAS-IkB tumor pool

GFP-RAS-IkB tumor pool

GFP-RAS-IkB tumor pool

Normal breast tissue

Supplementary Table 5 Primers and Oligos

RACE primers for PANDA

Fwd 5'-CAGAACTTGGCATGATGGAG-3'

Rev 5'-TGATATGAAACTCGGTTTACTACTAGC-3'

Fwd2 5'-TGCACACATTTAACCCGAAG-3'
Rev2 5'-CCCCAAAGCTACATCTATGACA-3'
Rev3 5'-CGTCTCCATCAT GCCAAGTT-3'
Rev4 5'-CATAGAGCTTCACCGACATAGC-3'

RT-PCR primers for PANDA

Fwd 5'-TGCACACATTTAACCCGAAG-3'
Rev 5'-CCCCAAAGCTACATCTATGACA-3'

siRNAs for PANDA

siRNA pool A 5'-AAUGUGUGCACGUAACAGAUU-3'

5'-GAGAUUUGCAGCAGACACAUU-3'

siRNA pool B 5'-GGGCAUGUUUUCACAGAGGUU-3'

5'-GAGAUUUGCAGCAGACACAUU-3'

siRNA pool C 5'-AAUGUGUGCACGUAACAGAUU-3'

5'-GGGCAUGUUUUCACAGAGGUU-3'

siCTRL Dharmacon D-001810-10

siRNAs for mRNAs

siNFYA pool si9530 Ambion

si9529 Ambion si9528 Ambion

siTP53 S606 Ambion siCDKN1A S417 Ambion

Chip primers

PUMA fwd 5'-CGT GGA TTC CTG TCT CCT CT-3'
PUMA rev 5'-GTC ACT CTG GTG AGG CGA TT-3'
NOXA fwd 5'-TTT CCC TTC CCT GTT ACT GC-3'
NOXA rev 5'-CTT GGG TAA ACA AGC CCA GA-3'

Taqman assays

 PANDA
 custom Taqman

 TP53
 Hs99999147_m1

 LAP3
 Rh02870758_m1

 APAF1
 Hs00559441_m1

 LRDD
 Hs00388035_m1

FAS	Hs00163653_m1
BIK	Hs00154189_m1
CDKN1A	Hs01121168_m1
GAPDH	Hs99999905 m1